

**FOOD AND DRUG ADMINISTRATION (FDA)
Center for Drug Evaluation and Research (CDER)**

Pediatric Subcommittee of the Oncologic Drugs Advisory Committee (pedsODAC) Meeting

FDA White Oak Campus, Building 31, The Great Room (Rm. 1503)

White Oak Conference Center, Silver Spring, Maryland

June 28 - 29, 2016

AGENDA

On June 28, 2016, information will be presented for expert assessments related to exploring potential pediatric development plans for three products in various stages of development for adult cancer indications. The subcommittee will consider and discuss issues concerning diseases to be studied, patient populations to be included, and possible study designs in the development of these products for pediatric use. The discussion will also provide information to the Agency pertinent to the formulation of written requests for pediatric studies, if appropriate. The products under consideration are: (1) venetoclax, presentation by AbbVie, Inc. (2) tazemetostat, presentation by Epizyme, Inc., and (3) atezolizumab, presentation by Roche/Genentech.

8:00 a.m.	Call to Order Introduction of Subcommittee	Alberto Pappo, MD Chairperson, Pediatric Subcommittee of the Oncologic Drugs Advisory Committee (ODAC)
8:10 a.m.	FDA Introductory Remarks/Presentation	Gregory Reaman, MD Associate Director for Oncology Sciences Office of Hematology and Oncology Products (OHOP), Office of New Drugs (OND) CDER, FDA
8:15 a.m.	Topic 1: ABT-199 (Venetoclax) – AbbVie, Inc.	
	Conflict of Interest Statement	Lauren Tesh, PharmD, BCPS
8:20 a.m.	INDUSTRY PRESENTATION	AbbVie, Inc.
	Venetoclax for the Treatment of Pediatric Patients with Relapsed/Refractory Cancers	Su Young Kim, MD, PhD Medical Director, Oncology Development AbbVie, Inc.
8:40 a.m.	Clarifying Questions from Subcommittee	
8:50 a.m.	OPEN PUBLIC HEARING	
9:10 a.m.	Questions to the Subcommittee and Subcommittee Discussion	
10:10 a.m.	BREAK	
10:25 a.m.	Topic 2: Tazemetostat – Epizyme, Inc.	
	Conflict of Interest Statement	Lauren Tesh, PharmD, BCPS

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10:30 a.m.	INDUSTRY PRESENTATION	Epizyme, Inc.
	Tazemetostat for the Treatment of Pediatric Subjects with Malignant Rhabdoid Tumors and Other INI1-Negative Tumors	Peter Ho, MD, PhD Chief Medical Officer Epizyme, Inc.
10:50 a.m.	Clarifying Questions from Subcommittee	
11:00 a.m.	OPEN PUBLIC HEARING	
11:20 p.m.	Questions to the Subcommittee and Subcommittee Discussion	
12:20 pm.	LUNCH	
1:20p.m.	Topic 3: Atezolizumab- Roche/Genentech	
	Conflict of Interest Statement	Lauren Tesh, PharmD, BCPS
1:25 p.m.	INDUSTRY PRESENTATION	Roche/Genentech
	Atezolizumab Oncology Development	Raphaël Rousseau, MD, PhD Global Head, Pediatric Oncology Drug Development Group Genentech, a member of the Roche Group
1:45 p.m.	Clarifying Questions from Subcommittee	
1:55 p.m.	OPEN PUBLIC HEARING	
2:15 p.m.	Questions to the Subcommittee and Subcommittee Discussion	
3:15 p.m.	ADJOURNMENT	

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AGENDA (cont.)

On June 29, 2016, during the morning session, information will be presented for expert assessments related to exploring potential pediatric development plans for two products in various stages of development for adult cancer indications. The subcommittee will consider and discuss issues concerning diseases to be studied, patient populations to be included, and possible study designs in the development of these products for pediatric use. The discussion will also provide information to the Agency pertinent to the formulation of written requests for pediatric studies, if appropriate. The products under consideration are: (1) LOXO-101, presentation by Loxo Oncology, Inc., and (2) entrectinib, presentation by Ignyta, Inc.

During the afternoon session, information will be presented on the current unmet clinical need in the nearly uniformly fatal brain tumor, diffuse intrinsic pontine glioma (DIPG) which occurs predominantly in the pediatric age group. The diagnosis of DIPG is typically based on characteristic radiographic and clinical features in lieu of brain biopsy, and histological confirmation. Recent data has demonstrated that the biology and pathophysiology of these tumors differ. There are no approved drugs for this disease. Clinical investigators seek to exploit precision medicine approaches to DIPG and use potentially predictive information from the genomic signature of tumors at either diagnosis or relapse. This information can be used to select specific molecularly targeted drugs based on the genetic aberrations of an individual patient's tumor. The Agency will seek the input of the subcommittee, including an assessment of benefit/risk given the potential for an adverse event associated with a surgical intervention in the brainstem.

8:00 a.m.	Call to Order Introduction of Subcommittee	Alberto Pappo, MD Chairperson, Pediatric Subcommittee of the Oncologic Drugs Advisory Committee (ODAC)
8:10 a.m.	FDA Introductory Remarks/Presentation	Gregory Reaman, MD Associate Director for Oncology Sciences, Office of Hematology and Oncology Products (OHOP), Office of New Drugs (OND) CDER, FDA
8:15 a.m.	Topic 1: LOXO-101- Loxo Oncology, Inc. Conflict of Interest Statement	Lauren Tesh, PharmD, BCPS Designated Federal Officer, ODAC
8:20 a.m.	INDUSTRY PRESENTATION Developing LOXO-101 in Children with NTRK Gene Fusion Cancers	Loxo Oncology, Inc. Josh Bilenker, MD Chief Executive Officer Loxo Oncology, Inc.
8:40 a.m.	Clarifying Questions from Subcommittee	
8:50 a.m.	OPEN PUBLIC HEARING	
9:10 a.m.	Questions to the Subcommittee and Subcommittee Discussion	

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10:10 a.m.	BREAK	
10:20 a.m.	Topic 2: Entrectinib – Ignyta, Inc.	
	Conflict of Interest Statement	Lauren Tesh, PharmD, BCPS
10:25 a.m.	INDUSTRY PRESENTATION	Ignyta, Inc.
	Entrectinib for the Treatment of Pediatric Cancers Harboring an Activating Alteration of <i>NTRK1/2/3</i> , <i>ROS1</i> , or <i>ALK</i>	Pratik S. Multani, MD, MS Chief Medical Officer at Ignyta, Inc.
10:45 a.m.	Clarifying Questions from Subcommittee	
10:55 a.m.	OPEN PUBLIC HEARING	
11:15 p.m.	Questions to the Subcommittee and Subcommittee Discussion	
12:15 pm.	LUNCH	
1:15 p.m.	Topic 3: Diffuse Intrinsic Pontine Glioma (DIPG)	
	Conflict of Interest Statement	Lauren Tesh, PharmD, BCPS
1:20 p.m.	FDA Introductory Remarks	Joohee Sul, PhD Medical Officer Division of Oncology Products II (DOPII) OHOP, OND, CDER, FDA
1:25 p.m.	FDA PRESENTATIONS	
	To Biopsy or Not to Biopsy – That is the Question.	Robert (Skip) Nelson, MD Deputy Director and Senior Pediatric Ethicist Office of Pediatric Therapeutics Office of the Commissioner, FDA
	Biopsy Risks for Investigational in vitro Diagnostic Devices	Jeffrey D. Seidman, MD Medical Officer/Pathologist Molecular Pathology and Cytology Branch Division of Molecular Genetics and Pathology Office of In Vitro Diagnostics and Radiological Health Center for Devices and Radiological Health FDA

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| 2:10 p.m. | SPEAKER PRESENTATION

Treatment Opportunities in Diffuse
Intrinsic Pontine Glioma (DIPG) | Mark W. Kieran, MD, PhD
Director, Pediatric Medical Neuro-Oncology
Dana-Farber Cancer Institute/Boston Children's
Hospital
Director, Pediatric Brain Tumor Clinic
Dana-Farber Cancer Institute/Boston Children's
Hospital
Associate Professor of Pediatrics
Harvard Medical School |
| 2:25 p.m. | GUEST SPEAKER PRESENTATIONS

DIPG: The Role of Neurosurgery | Jeffrey R. Leonard, MD
Pediatric Neurosurgeon
Chief of Neurosurgery
Nationwide Children's Hospital
Professor, Department of Neurological Surgery
The Ohio State University |
| | Surgical Experience with Biopsy of
Brainstem Tumors | Nalin Gupta, MD, PhD
Professor in Residence of Neurological
Surgery and Pediatrics
Dennis Bruce Dettmer Endowed Chair in
Pediatric Neurosurgery
Director, Pediatric Neurological Surgery
Program
Principal Investigator, Brain Tumor
Research Center
University of California, San Francisco |
| | Clarifying Questions from Subcommittee | |
| 2:50 p.m. | BREAK | |
| 3:05 p.m. | OPEN PUBLIC HEARING | |
| 3:25 p.m. | Questions to the Subcommittee and
Subcommittee Discussion | |
| 4:25p.m. | Closing Remarks | Gregory Reaman, MD |
| 4:30 p.m. | ADJOURNMENT | |